Serial No.: 09/601,997

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IN THE CLAIMS:

Please amend claims 58 and 59 and add claim 75 as follows. This listing of claims replaces all prior such listings of claims.

LISTING OF CLAIMS:

- 1-8. (Cancelled)
- 9. (Previously presented) The method of claim 58, wherein said function is a physiological function.
- 10. (Previously presented) The method of claim 58, wherein said function is enzyme activity.
- (Previously presented) The method of claim 58, wherein said function is 11. protein synthesis.
- 12. (Previously presented) The method of claim 58, wherein said function is expression of a biological factor.
- (Previously presented) The method of claim 58, wherein said function is a 13. regulatory effector function.
- (Previously presented) The method of claim 58, wherein said phenotypic 14. change is monitored directly.
 - 15-57. (Cancelled)
- (Currently amended) A high-throughput method of assigning a function associated with a product encoded by a sample nucleic acid sequence in a target nucleic acid molecule, said method comprising:
- a) without any intervening bacterial cloning steps and without any conformational modeling of mRNA transcribed from the sample nucleic acid sequence in the target nucleic acid molecule, delivering into, amplifying and expressing a plurality of members of an oligonucleotide family as individual transcription products in a plurality of recombinant nonbacterial host cells comprising the target nucleic acid molecule that comprises the sample nucleic acid sequence, whereby the method is high-throughput, wherein:

the oligonucleotide family comprises a plurality of nucleic acid molecules; each member of the oligonucleotide family encodes a transcription product comprising a sequence that is complementary to a sequence contained in the mRNA transcribed from the sample nucleic acid sequence in the target nucleic acid molecule;

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the plurality of members of the oligonucleotide family are introduced into expression vectors, which are introduced into the host cells, wherein each the expression vectors vector comprises double-stranded DNA, comprising:

> a sense strand and an antisense strand, wherein the sense strand encodes a transcription product that is complementary to a sequence contained in the mRNA transcribed from the sample nucleic acid sequence in the target nucleic acid molecule and binds to an mRNA sequence transcribed from the sample nucleic acid sequence in the target nucleic molecule so that expression of a product coded for by the sample nucleic acid sequence is inhibited; and

> means for determining directionality of expression so that the doublestranded DNA is ligated into the delivery vector in the correct orientation for expression, wherein the product coded for by the sample nucleic acid sequence is associated with at least one phenotypic property of a host cell containing the mRNA sequence; and wherein the expression vector is for expression in nonbacterial host cells;

the coding sequence for each individual transcription product encodes an antisense nucleic acid that binds to the mRNA transcribed from the sample nucleic acid sequence in the target nucleic acid molecule;

the sequences of the mRNA to which the transcription product of the family members are complemenary are distributed throughout the target nucleic acid molecule; and

expression of one or more of the individual transcription products, but not necessarily all, inhibits production of a product of the mRNA,

- b) in the resulting host cells, comparing the phenotypes of the resulting host cells to phenotypes of control cells to identify changes in phenotype to thereby assign a function associated with the product encoded by the sample nucleic acid sequence in the target nucleic acid molecule, wherein control cells are untransfected host cells, whereby changes in phenotype can be assigned by comparison of the transfected host cell, and the un-transfected host cell.
- 59. (Currently Amended) The method of claim 58, wherein the transcription product that is encoded by the sense strand and binds to an mRNA sequence transcribed from the sample nucleic acid sequence in the target nucleic acid molecule so that expression of a product encoded by the sample nucleic acid sequence is inhibited, comprises:

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> a catalytic domain that cleaves the mRNA sequence transcribed from the sample nucleic acid in the target nucleic molecule; and

binding sequences flanking the catalytic domain for binding the transcription product to the mRNA, and/or wherein the means for determining directionality of expression comprises a different non blunt-ended restriction enzyme site at each end of said double-stranded DNA.

- 60. (Original) The method of claim 59, wherein the double-stranded DNA is formed by contacting a first oligonucleotide with a complementary second oligonucleotide, and/or wherein the non blunt-ended restriction enzyme site is complementary to an end of the expression vector.
- 61. (Original) The method of claim 59, wherein said expression vector is formed by: (a) contacting a double-stranded oligonucleotide with an expression vector; or (b) by contacting a single-stranded oligonucleotide with said expression vector; or (c) contacting a triple-stranded oligonucleotide with an expression vector.
- 62. (Previously Presented) The method of claim 58, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.
- (Original) The method of claim 62, wherein the virus is a retrovirus or an 63. adeno-associated virus.
- 64. (Previously Presented) The method of claim 58, wherein the expression vector is transfected directly into mammalian cells.
- 65. (Previously Presented) The method of claim 58, wherein the sample nucleic acid is genomic DNA, cDNA, an expressed sequence tag (EST) or RNA.
- 66. (Previously Presented) The method of claim 58, wherein the family contains between 3 and 20 members.
- 67. (Previously presented) The method of claim 58, wherein each member of the family is designed to inhibit the production of a product of a sample nucleic acid sequence in the target nucleic acid molecule.
- (Previously Presented) The method of claim 58, whereby all members of a 68. family are assessed in a single experiment.
- (Previously presented) The method of claim 58, whereby a plurality of different target nucleic acid molecules comprising sample nucleic acid sequences are assessed.

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70. (Original) The method of claim 59, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.

- 71. (Original) The method of claim 60, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.
- 72. (Original) The method of claim 61, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.
- 73. (Previously Presented) The method of claim 58, wherein the oligonucleotide family is a ribozyme family.
 - 74. (Cancelled)
- (New) The method of claim 58, wherein the oligonucleotide family contains 75. about 20 members.